

(19) World Intellectual Property Organization
International Bureau



(43) International Publication Date
3 January 2003 (03.01.2003)

PCT

(10) International Publication Number
WO 03/000218 A2

(51) International Patent Classification⁷: **A61K 7/32, 7/48**

(21) International Application Number: **PCT/EP02/06376**

(22) International Filing Date: **10 June 2002 (10.06.2002)**

(25) Filing Language: **English**

(26) Publication Language: **English**

(30) Priority Data:
0115344.4 **22 June 2001 (22.06.2001)** **GB**

(71) Applicant (for AE, AG, AU, BB, BZ, CA, CY, GB, GD, GH, GM, IE, IL, KE, LC, LK, LS, MN, MW, NZ, OM, SD, SG, SL, SZ, TT, TZ, UG, ZA, ZM, ZW only): **UNILEVER PLC** [GB/GB]; Unilever House, Blackfriars, London EC4P 4BQ (GB).

(71) Applicant (for AL, AM, AT, AZ, BA, BE, BF, BG, BJ, BR, BY, CF, CG, CH, CI, CM, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, FR, GA, GE, GN, GQ, GR, GW, HR, HU, ID, IS, IT, JP, KG, KP, KR, KZ, LR, LT, LU, LV, MA, MC, MD, MG, MK, ML, MR, MX, MZ, NE, NL, NO, PH, PL, PT, RO, RU, SE, SI, SK, SN, TD, TG, TJ, TM, TN, TR, UA, UZ, VN, YU only): **UNILEVER NV** [NL/NL]; Weena 455, NL-3013 AL Rotterdam (NL).

(71) Applicant (for IN only): **HINDUSTAN LEVER LIMITED** [IN/IN]; Hindustan Lever House, 165/166 Backbay Reclamation, Maharashtra, 400 020 Mumbai (IN).

(72) Inventors; and

(75) Inventors/Applicants (for US only): **COX, Diana, Sheila**

[GB/GB]; Unilever R & D Colworth, Sharnbrook, Bedford, Bedfordshire MK44 1LQ (GB). **JAMES, Alexander, Gordon** [GB/GB]; Unilever R & D Colworth, Sharnbrook, Bedford, Bedfordshire MK44 1LQ (GB). **TAYLOR, David** [GB/GB]; Unilever Research Port Sunlight, Quarry Road East, Bebington, Wirral, Merseyside CH63 3JW (GB).

(74) Agents: **ELLIOTT, Peter, William et al.**; Unilever PLC, Patent Department, Colworth House, Sharnbrook, Bedford, Bedfordshire MK44 1LQ (GB).

(81) Designated States (*national*): AE, AG, AL, AM, AT (utility model), AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ (utility model), CZ, DE (utility model), DE, DK (utility model), DK, DM, DZ, EC, EE (utility model), EE, ES, FI (utility model), FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK (utility model), SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW.

(84) Designated States (*regional*): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

Published:

— without international search report and to be republished upon receipt of that report

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

WO 03/000218 A2

(54) Title: **COSMETIC COMPOSITIONS**

(57) Abstract: This invention concerns cosmetic compositions and methods involving 4-hydroxy-3-methoxybenzyl alcohol as an active ingredient. This material is shown to be an extremely effective sub-lethal inhibitor of the metabolism of selected corynebacteria and to give significant deodorancy benefits.

- 1 -

COSMETIC COMPOSITIONSField of Invention

- 5 This invention relates to cosmetic compositions for reducing or preventing body malodour. In particular, it relates to cosmetic compositions comprising a highly effective, sub-lethal inhibitor of selected corynebacteria.

10 Background

It is well known that freshly secreted sweat is sterile and that body malodour is the result of biotransformation of the sweat by microorganisms living on the surface of the skin to
15 produce volatile odoriferous compounds.

There are three types of material routinely used to combat body malodour: perfumes, antiperspirants and deodorants.

- 20 Perfumes typically work by simply masking body malodour.

Antiperspirants work by blocking the sweat glands, thereby reducing perspiration. However, even the best cosmetically acceptable antiperspirants rarely reduce sweat production by
25 more than 50%.

Typical deodorants work by reducing the population of microorganisms living on the surface of the skin, thereby reducing the extent of sweat biotransformation referred to
30 above. Typical deodorants include ethanol and triclosan (2,4,4'-trichloro,2'-hydroxy-diphenyl ether). However, the

- 2 -

skin is host to a number of species of microorganism, some of which are beneficial. The use of typical deodorants results in the killing of these beneficial species, in addition to the odour-producing species. This is an
5 undesirable side effect of such deodorants.

The present invention concerns malodour reduction via the sub-lethal inhibition of certain corynebacteria, as described in WO 00/01356 (Quest International BV) and WO
10 00/01353 (Unilever), the latter of which is incorporated herein by reference. These prior publications disclose the sub-lethal inhibition of corynebacteria that are capable of catabolising fatty acids. Many materials are described as having this effect; however, the highly effective
15 compositions of the present application are not disclosed.

WO 00/01353 (Unilever) uses the term "corynebacteria A" to mean corynebacteria that are able to catabolise fatty acids; this term is used with the same meaning in the present
20 application. Such bacteria contribute strongly to the formation of body malodour, in particular axillary malodour. For many males, malodour formation is largely caused by corynebacteria A.

25 The deodorants presently available on the market tend to be insufficiently effective or substantially reduce the numbers of all bacteria on the skin indiscriminately. The present invention offers the opportunity to provide cosmetic compositions, which, for many females, will substantially
30 reduce malodour formation while inactivating only a minor portion of the skin microflora. For many males, malodour

- 3 -

formation can be substantially reduced or even largely eliminated while inactivating only one subgroup of the skin microflora, the corynebacteria A.

- 5 Furthermore, the specific active ingredient disclosed in the present application is effective at particularly low concentrations.

Other publications in the prior art describe alternative
10 deodorancy methods that do not indiscriminately kill the skin microflora.

DD 29 39 58 (Medezinische Fakultaet [Charité] der Humboldt
Universitaet zu Berlin) describes the use of lipoxxygenase
15 inhibitors to act biochemically to reduce sweat production or to inhibit, to various degrees, the action of skin bacteria or their enzymes on the decomposition of sweat to form unpleasant-smelling substances.

20 DE 43 43 265 (Henkel) describes deodorant compositions comprising saturated dioic acid (C3-C10) esters. The active inhibits a sweat decomposing esterase and the compositions are said to not disturb the skin's natural microflora.

25 DE 43 43 264 (Henkel) describes the use of lipid-soluble partial esters of hydroxy carboxylic acids in deodorant compositions.

US 4,356,190 (Personal Products Co.) describes a deodorancy
30 method utilising selected aminopolycarboxylic acids that function whilst maintaining the viability of corynebacteria.

- 4 -

New deodorants containing p-hydroxybenzaldehyde or p-hydroxybenzyl alcohol are described in JP 63,292,962 (Matsushita Electric Works Ltd.).

5 Summary of the Invention

According to a first aspect of the invention, there is provided a cosmetic composition comprising 4-hydroxy-3-methoxybenzyl alcohol.

10

According to a second aspect of the invention, there is provided a cosmetic method of obtaining a deodorancy benefit comprising the topical administration of 4-hydroxy-3-methoxybenzyl alcohol.

15

Detailed Description

4-hydroxy-3-methoxybenzyl alcohol, the active ingredient utilised in the present invention, is capable of inhibiting fatty acid catabolism by corynebacteria A at a concentration below that which would lead to the death of said corynebacteria A. The active ingredient leads to a deodorancy benefit without significant harm to the skin's natural microflora. In addition, the active ingredient is effective at particularly low concentrations, being able to reduce the fatty acid catabolism of corynebacteria A by greater than 50% at a concentration of 0.5 mg/ml or less.

The aforementioned inhibitory effect may be described as sub-lethal, in that the effect is obtained at a concentration below that which would lead to the death of

30

- 5 -

the corynebacteria A. The effect may be further defined as a significant inhibition of fatty acid catabolism, for example greater than 50% inhibition of pentadecanoic acid utilisation, without a concomitant reduction in cell

5 viability ($\leq 1 \log_{10}$ CFU/ml reduction) of the corynebacteria A. The active ingredient is able to produce this effect at a concentration of 0.25 mg/ml or less.

The active ingredient may be employed in any cosmetic

10 composition. A particularly useful application is in deodorant compositions, particularly those used on the human body, and especially those used for treatment of underarm and/or foot malodour.

15 Compositions according to the invention comprise an effective total concentration of active ingredient; that is to say, a concentration sufficient to inhibit the catabolism of fatty acids by corynebacteria A on normal use of the composition. Typical concentrations range from 0.001 to

20 10%, preferably from 0.01 to 5%, and especially from 0.2 to 2% by weight of the composition.

In one aspect of the invention, it is desirable that the compositions of the invention do not comprise significant

25 amounts of additional anti-microbial agents that cause lethal inhibition of corynebacteria A. It is desirable that the total concentration of such anti-microbial agents is less than the total concentration of active ingredients according to the invention; indeed, it is preferred that the total

30 concentration of such anti-microbial agents is less than half, and especially less than one tenth, of this amount.

- 6 -

Active ingredients that cause lethal inhibition of corynebacteria A may be defined as those causing $\geq 1 \log_{10}$ CFU/ml reduction in cell viability when tested by methods common in the art, for example the method described in
5 Example 1 of the present specification (*vide infra*).

Cosmetic compositions according to the invention may take any of a variety of forms. Typical forms include aerosols, sticks, soft solids, creams, gels, roll-ons, pump sprays,
10 squeeze sprays, and compositions for application to deodorant wipes. All of the above forms are particularly applicable forms of deodorant composition.

Cosmetic compositions according to the invention comprise
15 one or more components in addition to the active ingredient. A commonly employed additional component is a carrier material. Such materials serve to aid the delivery of the active ingredient to the desired target. Preferred carrier materials are liquids at ambient temperature and atmospheric
20 pressure. Hydrophobic liquids suitable for use include liquid silicones, that is to say, liquid polyorganosiloxanes. Such materials may be cyclic or linear, examples include Dow Corning silicone fluids 344, 345, 244, 245, 246, 556, and the 200 series; Union Carbide
25 Corporation Silicones 7207 and 7158; and General Electric silicone SF1202. Alternatively, non-silicone hydrophobic liquids may be used. Such materials include mineral oils, hydrogenated polyisobutene, polydecene, paraffins, isoparaffins of at least 10 carbon atoms, and aliphatic or
30 aromatic ester oils (e.g. isopropyl myristate, lauryl

- 7 -

myristate, isopropyl palmitate, diisopropyl sebecate, diisopropyl adipate, or C₈ to C₁₈ alkyl benzoates).

Hydrophilic liquid carrier materials, for example water, may
5 also be employed.

Particularly preferred liquid carrier materials comprise organic solvents. Preferred organic solvents have a melting point of less than 10°C, preferably less than 5°C; this can
10 benefit both low temperature storage stability and ease of manufacture. A class of preferred organic solvents are aliphatic alcohols (monohydric or polyhydric, preferably having 2 to 8 carbon atoms) and polyglycol ethers, preferably oligoglycol ethers having only 2 to 5 repeat
15 units. Examples include dipropylene glycol, glycerol propylene glycol, butylene glycol, ethanol, propanol, isopropanol, and industrial methylated spirits. The most preferred organic solvents are aliphatic alcohols, in particular those having 2 to 3 carbon atoms, especially
20 ethanol and isopropanol.

Mixtures of carrier materials may also be used. The total amount of carrier material employed is preferably from 1 to 99%, more preferably from 10% to 98%, and most preferably
25 from 50% to 97% by weight of the composition, excluding any volatile propellant that might also be present.

A variety of other materials may also be employed in the compositions of the invention. In certain aspects of the
30 invention, an additional deodorant active may be desirable.

- 8 -

This might be a perfume, an antiperspirant active, or an anti-microbial active.

Perfumes, when employed, may be conventional perfumes, such as perfume oils, and/or so-called deo-perfumes, as described in EP 545,556 and other publications. Levels of incorporation are preferably up to 4% by weight, particularly from 0.1% to 2% by weight, and especially from 0.7% to 1.7% by weight of the composition.

10

Compositions according to the invention that additionally comprise an antiperspirant active are particularly preferred. Typical antiperspirant actives include astringent active salts, in particular, aluminium, zirconium and mixed aluminium/zirconium salts, including both inorganic salts, salts with organic anions and complexes. Preferred astringent salts include aluminium, zirconium and aluminium/zirconium halides and halohydrate salts, such as chlorohydrates. Preferred levels of incorporation are from 0.5% to 60%, particularly from 5% to 30% or 40% and especially from 5% or 10% to 30% or 35% by weight of the composition of which it is a part. In non-aqueous formulations, the above weight percentages exclude any water of hydration bound to the antiperspirant salt. Especially preferred aluminium halohydrate salts, known as activated aluminium chlorohydrates, are described in EP 6,739 (Unilever PLC and NV). Zirconium aluminium chlorohydrate actives are also preferred materials, as are the so-called ZAG (zirconium-aluminium-glycine) complexes, for example those disclosed in US 3,792,068 (Procter and Gamble Co.).

30

- 9 -

Typical anti-microbial actives include quaternary ammonium compounds (like cetyltrimethylammonium salts), chlorhexidine and salts thereof; diglycerol monocaprates, diglycerol monolaurate, glycerol monolaurate, polyhexamethylene biguanide salts (also known as polyaminopropyl biguanide salts - an example being Cosmocil CQ available from Zeneca PLC), 2,4,4'-trichloro,2'-hydroxy-diphenyl ether (triclosan), and 3,7,11-trimethyldodeca-2,6,10-trienol (farnesol). Typical levels of incorporation are from 0.01% to 1%, in particular from 0.03% to 0.5%, or especially from 0.05% to 0.3% by weight of the composition.

Particularly preferred additional deodorant actives are agents that are capable of sub-lethal inhibition of corynebacteria A, in particular, sub-lethal inhibition of fatty acid catabolism by corynebacteria A. The effect may be further defined as a significant inhibition of fatty acid catabolism, for example greater than 50% inhibition of pentadecanoic acid utilisation, without a concomitant reduction in cell viability ($\leq 1 \log_{10}$ CFU/ml reduction) of the corynebacteria A. Such agents may be used in concentrations ranging from 0.001 to 10%, in particular from 0.05 to 5%, and especially from 0.3 to 3% by weight of the composition. Examples of such agents are described in WO 00/01356 (Quest International BV) and WO 00/01353 (Unilever). Other examples are the chelating agents described in US 4,356,190 (Personal Products Co.) and/or our co-pending application PCT/EP01/00118 (Unilever), particularly those chelating agents having an iron (III) binding constant of greater than 10^{26} . DTPA

- 10 -

(diethylenetriaminepentaacetic acid) and salts thereof are especially preferred.

Structurants and emulsifiers are further additional
5 components of the compositions of the invention that are highly desirable in certain product forms. Structurants, when employed, are preferably present at from 1% to 30% by weight of the composition, whilst emulsifiers are preferably present at from 0.1% to 10% by weight of the composition.
10 Suitable structurants include cellulosic thickeners such as hydroxy propyl cellulose and hydroxy ethyl cellulose, and dibenzylidene sorbitol. Emulsion pump sprays, roll-ons, creams, and gel compositions according to the invention can be formed using a range of oils, waxes, and emulsifiers.
15 Suitable emulsifiers include steareth-2, steareth-20, steareth-21, cetareth-20, glyceryl stearate, cetyl alcohol, cetearyl alcohol, PEG-20 stearate, and dimethicone copolyol. Suspension aerosols, roll-ons, sticks, and creams require structurants to slow sedimentation (in fluid compositions)
20 and to give the desired product consistency to non-fluid compositions. Suitable structurants include sodium stearate, stearyl alcohol, cetyl alcohol, hydrogenated castor oil, synthetic waxes, paraffin waxes, hydroxystearic acid, dibutyl lauroyl glutamide, alkyl silicone waxes,
25 quaternium-18 bentonite, quaternium-18 hectorite, silica, and propylene carbonate. Some of the above materials also function as suspending agents in certain compositions.

Further emulsifiers desirable in certain compositions of the
30 invention are perfume solubilisers and wash-off agents. Examples of the former include PEG-hydrogenated castor oil,

- 11 -

available from BASF in the Cremaphor RH and CO ranges, preferably present at up to 1.5% by weight, more preferably 0.3 to 0.7% by weight. Examples of the latter include poly(oxyethylene) ethers.

5

Sensory modifiers are further desirable components in certain compositions of the invention. Such materials are preferably used at a level of up to 20% by weight of the composition. Emollients, humectants, volatile oils, non-
10 volatile oils, and particulate solids which impart lubricity are all suitable classes of sensory modifiers. Examples of such materials include cyclomethicone, dimethicone, dimethiconol, isopropyl myristate, isopropyl palmitate, talc, finely-divided silica (e.g. Aerosil 200), polyethylene
15 (eg. Acumist B18), polysaccharides, corn starch, C12-C15 alcohol benzoate, PPG-3 myristyl ether, octyl dodecanol, C7-C14 isoparaffins, di-isopropyl adipate, isosorbide laurate, PPG-14 butyl ether, glycerol, hydrogenated polyisobutene, polydecene, titanium dioxide, phenyl trimethicone, dioctyl
20 adipate, and hexamethyl disiloxane.

Cosmetic compositions that are aerosols generally also comprise a volatile propellant. The propellant may be selected from liquified hydrocarbons or halogenated
25 hydrocarbon gases (particularly fluorinated hydrocarbons such as 1,1-difluoroethane and/or 1-trifluoro-2-fluoroethane) that have a boiling point of below 10°C and especially those with a boiling point below 0°C. It is especially preferred to employ liquified hydrocarbon gases,
30 and especially C₃ to C₆ hydrocarbons, including propane,

- 12 -

isopropane, butane, isobutane, pentane and isopentane and mixtures of two or more thereof. Preferred propellants are isobutane, isobutane/isopropane, isobutane/propane and mixtures of isopropane, isobutane and butane.

5

Other propellants that can be contemplated include alkyl ethers, such as dimethyl ether or compressed non-reactive gasses such air, nitrogen or carbon dioxide.

- 10 Other additional components that may also be included are colourants and preservatives, for example C₁-C₃ alkyl parabens.

Examples

15

- This experiment uses the methods of WO 00/01353 to illustrate the much greater efficacy of 4-hydroxy-3-methoxybenzyl alcohol at inhibiting fatty acid catabolism by corynebacteria A, when compared with analogous materials previously disclosed in the aforementioned publication. Data illustrating the sub-lethal aspect of the inhibition are also presented.
- 20

- An *in vitro* model system, reproducing fatty acid catabolism by axillary bacteria, was used. To each of several 250 ml baffled shake flasks was added 30 ml semi-synthetic medium (see below), supplemented with fatty acid substrate (2.0 mg/ml pentadecanoic acid) and non-fatty acid substrate (0.5 mg/ml glucose). To each flask (other than the control) was also added one of the indicated test materials, as a 10 % (w/v) emulsion in semi-synthetic medium, supplemented with
- 25
- 30

- 13 -

Gum Arabic (5.0 mg/ml). (Emulsions were formed by ultra-homogenisation at 24,000 rpm for about 1 min.) Each of the flasks was inoculated with fresh bacterial biomass (*Corynebacterium* A sp. NCIMB 40928), pre-grown for 24 h in
5 TSBT (see below), to give starting optical densities (A_{590}) of 1.0-2.0. Following inoculation, the flasks were incubated aerobically at 35°C, with agitation (130 rpm), for 24 hours. After this time, culture viability and remaining fatty acid were determined by methodology as described in WO 99/01359.

10

Composition of semi-synthetic medium in g/l: KH_2PO_4 (1.6), $(\text{NH}_4)_2\text{HPO}_4$ (5.0), Na_2SO_4 (0.38), yeast nitrogen base (3.35) (Difco), yeast extract (0.5) (Beta Lab), Tween 80TM (0.2), Triton X-100TM (0.2), and $\text{MgCl}_2 \cdot 6\text{H}_2\text{O}$ (0.5).

15

Composition of TSBT (Tween-supplemented Tryptone soya broth) in g/l: Tryptone soya broth (30.0) (Merck), yeast extract (10.0) (Beta Lab), and Tween 80TM (1.0).

20 Table 1 illustrates the effects of the indicated actives upon *Corynebacterium* A sp. NCIMB 40928 in terms of culture viability and fatty acid utilisation. Various concentrations of active were investigated. It will be noted that for each of the Examples, culture viability was not substantially
25 affected.

- 14 -

Table 1: Effect of Actives upon *Corynebacterium* A sp. NCIMB 40928 (Comparative examples are indicated by letter codes)

Example	Conc. (g/l)	Viability (log ₁₀ CFU/ml)	Fatty acid utilisation (%)
1.	0	9.12	100
4-hydroxy-3-methoxybenzyl	0.1	8.91	65
alcohol	0.25	8.88	40
	0.5	8.28	20
A.	0	8.17	100
4-hydroxybenzyl	0.1	8.69	100
alcohol	0.25	8.85	97
	0.5	8.17	21
B.	0	8.98	100
3-hydroxy-4-methoxybenzyl	1.0	8.90	99
alcohol	5.0	8.99	86
	10.0	8.62	29
C.	0	8.75	100
4-hydroxy-3-methoxyphenethyl	1.0	8.80	99
alcohol	5.0	8.60	59
	10.0	8.09	26

- 5 These data illustrate that the active of the invention is an effective inhibitor of fatty acid catabolism by corynebacteria A at a considerably lower concentration than analogous materials disclosed in the prior art.

- 15 -

Examples 3 to 11

The following are typical compositions according to the invention and were prepared by methods common in the art.

- 5 Examples 3 to 8 are aerosol compositions, Example 9 is a pump spray composition, Example 10 is an antiperspirant stick composition, and Example 11 is a roll-on composition.

Table 2: Composition of Examples 3 to 8

10 (amounts given in the Tables are percentages by weight)

Example:	3	4	5	6	7	8
CAP 40 ¹	92	85	35	84.96	85	35
Ethanol (96%)	0	0	62.17	0	0	61.16
DC 245 ²	6.2	6.9	0	6.4	6.5	0
AACH ³		5	0	5	5	0
4-hydroxy-3-methoxybenzyl alcohol	1.0	2.0	1.0	1.0	1.0	1.0
Bentone 38 ⁴	0.6	0.5	0	0.5	0.5	0
DTPA ⁵	0	0	0	1.0	0	0
Cosmocil stearate ⁶	0	0	0	0.04	0	0
Irgasan DP-300 ⁷	0	0	0	0	0	0.01
Ferulic acid ⁸	0	0	0	0	1.0	1.0
Perfume	0	0.6	1.5	1.0	1.0	1.5
Isopropyl myristate	0	0	0.33	0	0	0.33
Propylene carbonate	0.2	0	0	0	0	0

1. Mixture of butane, isobutane and propane, ex Calor.

- 16 -

2. Cyclomethicone, ex Dow Corning.
3. Activated aluminium chlorohydrate, grade A296, ex Giulini.
4. Quaternium-18 hectorite, ex Rheox.
- 5 5. Diethylenetriaminepentaacetic acid, sieved to <63 µm.
6. Polyhexamethylene biguanide salt, ex Zeneca.
7. Triclosan, ex Ciba-Geigy.
8. 4-Hydroxy-3-methoxycinnamic acid, a deodorant active as disclosed in WO 00/01359 (Unilever).

10

Example 4 was found to have a significantly better deodorancy performance than a control composition having the 4-hydroxy-3-methoxybenzyl alcohol replaced by DC 245. A similar benefit was obtained with an analogous composition

15 comprising only 1% (w/w) 4-hydroxy-3-methoxybenzyl alcohol.

- 17 -

Table 3: Composition of Examples 9, 10, and 11

Example:	9	10	11
Ethanol	59.4	0	70
Water	39.6	0	27.85
4-hydroxy-3-methoxybenzyl alcohol	1.0	1.0	1.0
Cremaphor RH40 ¹	0	0	0.5
Klucel M ²	0	0	0.65
AAZG ³	0	25	0
DC 245	0	50.8	0
Stearyl alcohol	0	14	0
Superfino talc	0	3.2	0
PEG-8 distearate	0	1	0
Castorwax MP80	0	4	0
Perfume	0	1	0

1. PEG-hydrogenated castor oil, ex BASF.
2. Hydroxypropylcellulose, ex Aqualon.
- 5 3. Aluminium zirconium tetrachlorohydrex-glycine, Q5-7167,
ex Summit.

Example 10 was found to have a significantly better deodorancy performance than a control composition having the

10 4-hydroxy-3-methoxybenzyl alcohol replaced by DC 245. A similar benefit was obtained with an analogous composition comprising 2% (w/w) 4-hydroxy-3-methoxybenzyl alcohol and 49.8% (w/w) DC 245.

- 18 -

Examples 12 to 17

Tables 4 to 9 illustrate other compositions according to the invention that may be prepared by methods common in the art.

Table 4: Composition of Examples 12.1 to 12.6 (aerosol compositions)

Example:	12.1	12.2	12.3	12.4	12.5	12.6
Cyclomethicone (DC 245)	3.47	11.8	14.4	3.55	4.1	5.2
Ethanol			20			
Isopropyl palmitate			10.3		8.5	
Isopropyl myristate						0.31
PPG-14 butyl ether	9.7	0.7				9.1
Octyldodecanol		0.25				
Polydecene						0.3
Dibutyl phthalate					4.5	
Bentone 38 (ex Rheox)	1	1	1.5	1	0.95	0.7
Propylene carbonate					0.15	
Methylpropanolamine						0.08
Silicone gum (Q2-1401)				0.2		
AACH		10		4		
Milled AACH	10					2
Aluminium chlorohydrate			9.2		9.3	
Silica		0.1				0.01
Talc			3			
Micronised polyethylene					9.3	
Perfume	0.5	0.7	0.7	0.7		1
Allantoin					1.5	
Palmitoyl ethanolamide	0.3	0.3	0.3	0.3	0.3	0.3
4-hydroxy-3-methoxybenzyl alcohol	0.03	0.15	0.6	0.25	1.4	1
n-Pentane				20		
C3/C4 hydrocarbons	75	75	40	70	60	80

Table 5: Composition of Examples 13.1 to 13.9 (lotion compositions)

[illegible]

- 21 -

Table 6: Composition of Examples 14.1 to 14.5 (cream and soft solid compositions)

Example:	14.1	14.2	14.3	14.4	14.5
C18-C36 acid glycol ester		2.5		3.75	
Castor wax		7.5		1.25	
Triacontenyl vinyl pyrrolidone copolymer	5				
Paraffin wax	5				
Silica		1			0.2
Cyclopentasiloxane and cetearyl-dimethicone/vinyl dimethicone co-polymer					64.05
C12-15 alkyl benzoate	64.3	63.1	62.9	63.7	4
Dextrin palmitate			10	5	
Neopentyl glycol diheptanoate					5
PEG-8 distearate					2
Stearyl dimethicone					0.75
AACH	25			25.5	
Milled AACH		25.5	26		
AAZG					22
4-hydroxy-3-methoxybenzyl alcohol	0.2	0.4	0.6	0.8	1.5
Perfume	0.5		0.5		0.5

- 22 -

Table 7: Composition of Examples 15.1 to 15.8 (further cream and soft solid compositions)

Examples:	15.1	15.2	15.3	15.4	15.5	15.6	15.7	15.8
Silicone wax	2.5			3				
N-lauroyl glutamic acid dibutylamide		1						
C18-C36 acid glycol ester			5					
C18-C36 acid triglyceride			1.25					
Castor wax						4		
Stearyl alcohol						6		
Paraffin wax	7.5							
Candelilla wax							7	
C24/28 alkyl dimethicone wax							3.5	
Silica				1.5	1.5			
Talc			1.75		6	5		
Bentone 38					3		0.5	
Anhydrous aluminium silicate					6			
Microthene powder					6			
Propylene carbonate					1.5			
Cyclomethicone	64.4		61	62.5	36.3	56	43	47.8
Tetraphenyl tetramethylsiloxane		52.7						
C12-15 Alkyl benzoate				10				11.7
Dextrin palmitate		5						9

- 23 -

Octyldodecanol		15						
PPG14 butyl ether						4.5		
Dimethicone (10 mPa.s)			5		10			
Dimethicone (350 mPa.s)							24	
POE-100 stearyl ether					2			
POE-100 stearate						1		
AACH	25.5			22				
Milled AACH		25.5						
Aluminium chlorohydrate							18	
AAZG			25		25.7	20		26.5
4-hydroxy-3-methoxybenzyl alcohol	0.1	0.3	0.5	1	2	3	4	5
Perfume		0.5	0.5			0.5		

Table 8: Composition of Examples 16.1 to 16.6 (solid stick compositions)

Examples:	16.1	16.2	16.3	16.4	16.5	16.6
Cyclomethicone (DC245)	40.7	37.3	40.1	39.75	45.5	
Permethyl 103A	16	12				
PPG-14 Butyl ether		4	10			
Propylene glycol						47.8
Ethanol						13
Isostearyl alcohol						12
Stearyl alcohol	14	14	17	11.5		
Castor wax	2	5	2.5	5		
12-hydroxystearic acid					6	
N-lauroyl glutamic acid dibutylamide					2	
Dibenzylidene sorbitol						3
Eicosanol	0.2	0.2				
Octyldodecanol				14	14	
C20-40 alcohols					0.5	
C20-40 pareth-3/C20-40 pareth- 20				1.75		
PEG-8 distearate			0.6		5	
Amino-2-methyl-1-propanol						0.2
ZAG	23	25	24	26	26	22.5
Glycerol			2			
EDTA				1		
Talc	3					
Fumed silica		1	2			
Perfume	1	1	1			
4-hydroxy-3-methoxybenzyl alcohol	0.1	0.5	0.8	1	1	1.5

Table 9: Composition of Examples 17.1 to 17.6 (further solid stick compositions)

Examples:	17.1	17.2	17.3	17.4	17.5	17.6
Cyclomethicone (DC245)	36.3	49.25	10	37		
Mineral oil	11.5					
Polydecene			12.7			
PPG-14 butyl ether			2.5			
C12-15 alkyl benzoate				15		
Dimethicone (50 mPa.s)	1.5					
Propylene glycol					31	53.5
Ethanol					50	
Water					8.7	20
Stearyl alcohol	14				1	
Castor wax	4.5					
Dextrin palmitate		10				
Cellobiose octanonanoate			3.8			
Beta sitosterol				2.5		
Oryzanol				2.5		
Sodium stearate					5.8	7.7
Eicosanol	0.2					
Isopropyl myristate		10				
Cetyl dimethicone copolyol			1	1		
Amino-2-methyl-1-						0.5

- 26 -

propanol						
Poloxamer 407						6
Cocamide DEA						7
Aluminium chlorohydrate	26	30				
Zirkonal 50			51.7	40		
Triclosan						0.3
Glycerol	2		17.3			
Talc	1.5					
Fumed silica	1					
Perfume	1					
4-hydroxy-3- methoxybenzyl alcohol	0.5	0.75	1	2	3.5	5

- 27 -

CLAIMS:

1. A cosmetic composition comprising 4-hydroxy-3-methoxybenzyl alcohol.
5
2. A cosmetic composition according to claim 1, also comprising a carrier material.
3. A cosmetic composition according to claim 1 or claim 2,
10 wherein the 4-hydroxy-3-methoxybenzyl alcohol is present at a concentration of from 0.001 to 10% by weight.
4. A cosmetic composition according to claim 3, wherein
15 the 4-hydroxy-3-methoxybenzyl alcohol is present at a concentration of from 0.2 to 2% by weight.
5. A cosmetic composition according to any of the preceding claims, comprising an additional deodorant
20 active.
6. A cosmetic composition according to claim 5, wherein the additional deodorant active is capable of sub-lethal inhibition of corynebacteria A.
25
7. A cosmetic composition according to claim 5, wherein the additional deodorant active is an antiperspirant active.
- 30 8. A cosmetic composition according to any of claims 1 to 6, wherein said composition does not comprise

- 28 -

significant amounts of additional anti-microbial agents that cause lethal inhibition of corynebacteria A.

9. A cosmetic method of obtaining a deodorancy benefit
5 comprising the topical administration of 4-hydroxy-3-methoxybenzyl alcohol.
10. A cosmetic method of obtaining a deodorancy benefit
comprising the topical administration of a composition
10 according to any of claims 1 to 8.

(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization
International Bureau



(43) International Publication Date
3 January 2003 (03.01.2003)

PCT

(10) International Publication Number
WO 03/000218 A3

(51) International Patent Classification⁷: **A61K 7/32, 7/48**

(21) International Application Number: **PCT/EP02/06376**

(22) International Filing Date: **10 June 2002 (10.06.2002)**

(25) Filing Language: **English**

(26) Publication Language: **English**

(30) Priority Data:
0115344.4 22 June 2001 (22.06.2001) GB

(71) Applicant (for AE, AG, AU, BB, BZ, CA, CY, GB, GD, GH, GM, IE, IL, KE, LC, LK, LS, MN, MW, NZ, OM, SD, SG, SL, SZ, TT, TZ, UG, ZA, ZM, ZW only): **UNILEVER PLC** [GB/GB]; Unilever House, Blackfriars, London EC4P 4BQ (GB).

(71) Applicant (for AL, AM, AT, AZ, BA, BE, BF, BG, BJ, BR, BY, CF, CG, CH, CI, CM, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, FR, GA, GE, GN, GQ, GR, GW, HR, HU, ID, IS, IT, JP, KG, KP, KR, KZ, LR, LT, LU, LV, MA, MC, MD, MG, MK, ML, MR, MX, MZ, NE, NL, NO, PH, PL, PT, RO, RU, SE, SI, SK, SN, TD, TG, TJ, TM, TN, TR, UA, UZ, VN, YU only): **UNILEVER NV** [NL/NL]; Weena 455, NL-3013 AL Rotterdam (NL).

(71) Applicant (for IN only): **HINDUSTAN LEVER LIMITED** [IN/IN]; Hindustan Lever House, 165/166 Backbay Reclamation, Maharashtra, 400 020 Mumbai (IN).

(72) Inventors; and

(75) Inventors/Applicants (for US only): **COX, Diana, Sheila** [GB/GB]; Unilever R & D Colworth, Sharnbrook, Bedford, Bedfordshire MK44 1LQ (GB). **JAMES, Alexander,**

Gordon [GB/GB]; Unilever R & D Colworth, Sharnbrook, Bedford, Bedfordshire MK44 1LQ (GB). **TAYLOR, David** [GB/GB]; Unilever Research Port Sunlight, Quarry Road East, Bebington, Wirral, Merseyside CH63 3JW (GB).

(74) Agents: **ELLIOTT, Peter, William** et al.; Unilever PLC, Patent Department, Colworth House, Sharnbrook, Bedford, Bedfordshire MK44 1LQ (GB).

(81) Designated States (national): AE, AG, AL, AM, AT (utility model), AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ (utility model), CZ, DE (utility model), DE, DK (utility model), DK, DM, DZ, EC, EE (utility model), EE, ES, FI (utility model), FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK (utility model), SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW.

(84) Designated States (regional): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

Published:

— with international search report

(88) Date of publication of the international search report:
25 September 2003

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

WO 03/000218 A3

(54) Title: **COSMETIC COMPOSITIONS**

(57) Abstract: This invention concerns cosmetic compositions and methods involving 4-hydroxy-3-methoxybenzyl alcohol as an active ingredient. This material is shown to be an extremely effective sub-lethal inhibitor of the metabolism of selected corynebacteria and to give significant deodorancy benefits.

INTERNATIONAL SEARCH REPORT

Internat. Application No.

PCT/EP 02/06376

A. CLASSIFICATION OF SUBJECT MATTER
 IPC 7 A61K7/32 A61K7/48

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 A61K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

WPI Data, EPO-Internal, PAJ, CHEM ABS Data

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	DATABASE WPI Week 198413 Derwent Publications Ltd., London, GB; AN 1984-078371 XP002227570 & JP 59 029619 A (HASEGAWA CO), 16 February 1984 (1984-02-16) abstract	1,2
X	PATENT ABSTRACTS OF JAPAN vol. 17, no. 482 (C-1105), 2 September 1993 (1993-09-02) & JP 05 117972 A (SHINKI SANGYO K.K.), 14 May 1993 (1993-05-14) abstract	1
	-/--	

☒ Further documents are listed in the continuation of box C.

☐ Patent family members are listed in annex.

* Special categories of cited documents:

- *A* document defining the general state of the art which is not considered to be of particular relevance
- *E* earlier document but published on or after the International filing date
- *L* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- *O* document referring to an oral disclosure, use, exhibition or other means
- *P* document published prior to the international filing date but later than the priority date claimed

- *T* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- *X* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- *Y* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
- *G* document member of the same patent family

Date of the actual completion of the international search

16 January 2003

Date of mailing of the international search report

30/01/2003

Name and mailing address of the ISA

European Patent Office, P.B. 5818 Patentlaan 2
 NL - 2280 HV Rijswijk
 Tel. (+31-70) 340-2040. Tx. 31 651 epo nl.
 Fax: (+31-70) 340-3016

Authorized officer

Willekens, G

INTERNATIONAL SEARCH REPORT

Intern I Application No

PCT/EP 02/06376

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	DATABASE WPI Week 200027 Derwent Publications Ltd., London, GB; AN 2000-313125 XP002227571 & JP 2000 096078 A (HASEGAWA CO ET AL.), 4 April 2000 (2000-04-04) abstract ---	1,9
A	PATENT ABSTRACTS OF JAPAN vol. 2000, no. 10, 17 November 2000 (2000-11-17) & JP 2000 191520 A (SAKUMA KAZUO), 11 July 2000 (2000-07-11) abstract ---	1
A	DATABASE WPI Week 198511 Derwent Publications Ltd., London, GB; AN 1985-066384 XP002227572 & JP 60 023309 A (LION CORP), 5 February 1985 (1985-02-05) abstract ---	1
A	PATENT ABSTRACTS OF JAPAN vol. 1999, no. 9, 30 July 1999 (1999-07-30) & JP 11 116460 A (KAO CORP), 27 April 1999 (1999-04-27) abstract -----	1

INTERNATIONAL SEARCH REPORT

Information on patent family members

Internat Application No

PCT/EP 02/06376

Patent document cited in search report		Publication date	Patent family member(s)	Publication date
JP 59029619	A	16-02-1984	JP 1612334 C JP 2034322 B	30-07-1991 02-08-1990
JP 05117972	A	14-05-1993	NONE	
JP 2000096078	A	04-04-2000	JP 3304219 B2 JP 7179328 A	22-07-2002 18-07-1995
JP 2000191520	A	11-07-2000	NONE	
JP 60023309	A	05-02-1985	JP 62040328 B	27-08-1987
JP 11116460 0	A		NONE	

THIS PAGE BLANK (USPTO)